

FIG. 1A

KCNQ5 cDNA coding sequence

atgaaggatg tggagtcggg ccggggcagg gtgctgctga 40
actcggcagc cgccaggggc gacggcctgc tactgctggg 80
caccgcgcg gccacgcttg gtggcggcgg cggtggcctg 120
agggagagcc gccggggcaa gcagggggcc cggatgagcc 160
tgctggggaa gccgctctct tacacgagta gccagagctg 200
ccggcgcaac gtcaagtacc ggcgggtgca gaactacctg 240
tacaacgtgc tggagagacc ccgcggctgg gcgttcatct 280
accacgcttt cgtttttctc cttgtctttg gttgcttgat 320
tttgtcagtg ttttctacca tccctgagca cacaaaattg 360
gcctcaagtt gcctcttgat cctggagttc gtgatgattg 400
tcgtctttgg tttggagttc atcattcgaa tctggtctgc 440
gggttgctgt tgtcgatata gaggatggca aggaagactg 480
aggtttgctc gaaagccctt ctgtgttata gataccattg 520
ttcttategc ttcaatagca gttgtttctg caaaaactca 560
gggtaatatt tttgccacgt ctgcactcag aagtctccgt 600
ttcctacaga tcctccgcac ggtgcgcatg gaccgaaggg 640
gaggcacttg gaaattactg gggttcagtgg tttatgctca 680
cagcaaggaa ttaatcacag cttggtacat aggatttttg 720
gttcttattt tttcgtcttt ccttgtctat ctggtggaaa 760

FIG. 1B

aggatgccaa taaagagttt tctacatatg cagatgctct 800
ctggtggggc acaattacat tgacaactat tggctatgga 840
gacaaaactc ccctaacttg gctgggaaga ttgctttctg 880
caggctttgc actccttggc atttctttct ttgcacttcc 920
tgccggcatt cttggctcag gttttgcatt aaaagtacaa 960
gaacaacacc gccagaaaca ctttgagaaa agaaggaacc 1000
cagctgccaa cctcattcag tgtgtttggc gtagttacgc 1040
agctgatgag aaatctgttt ccattgcaac ctggaagcca 1080
cacttgaagg ccttgcacac ctgcagccct accaagaaag 1120
aacaagggga agcatcaagc agtcagaagc taagttttaa 1160
ggagcgagtg cgcattggcta gcccagggg ccagagtatt 1200
aagagccgac aagcctcagt aggtgacagg aggtccccaa 1240
gcaccgacat cacagccgag ggagtcacca ccaaagtgca 1280
gaagagctgg agcttcaacg accgaaccg cttccggccc 1320
tcgctgcgcc tcaaaagttc tcagccaaaa ccagtgatag 1360
atgctgacac agcccttggc actgatgatg tatatgatga 1400
aaaaggatgc cagtgtgatg tatcagtgga agacctcacc 1440
ccaccactta aaactgtcat tcgagctatc agaattatga 1480
aatttcatgt tgcaaaacgg aagtttaagg aaacgttacg 1520
tccatatgat gtaaaagatg tcattgaaca atattctgct 1560

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FIG. 1C

ggatcatctgg acatgtttgtg tagaattaaa agccttcaaa 1600
 cacgtgttga tcaaattctt ggaaaagggc aaatcacatc 1640
 agataagaag agccgagaga aaataacagc agaacatgag 1680
 accacagacg atctcagtat gctcggtcgg gtggtcaagg 1720
 ttgaaaaaca ggtacagtcc atagagtcca agctggactg 1760
 cctactagac atctatcaac aggtccttcg gaaaggctct 1800
 gcctcagccc tcgctttggc ttcattccag atcccacctt 1840
 ttgaatgtga acagacatct gactatcaaa gcctgtgga 1880
 tagcaaagat ctttcgggtt ccgcacaaaa cagtggctgc 1920
 ttatccagat caactagtgc caacatctcg agaggcctgc 1960
 agttcattct gacgccaaat gagttcagtg cccagacttt 2000
 ctacgcgctt agccctacta tgcacagtca agcaacacag 2040
 gtgccaatta gtcaaagcga tggctcagca gtggcagcca 2080
 ccaacaccat tgcaaacc aaataacagg caccacagcc 2120
 agcagcccca acaactttac agatcccacc tcctctccca 2160
 gccatcaagc atctgcccag gccagaaact ctgcacccta 2200
 accctgcagg cttacaggaa agcatttctg acgtcaccac 2240
 ctgccttggt gcctccaagg aaaatgttca ggttgcacag 2280
 tcaaattctca ccaaggaccg ttctatgagg aaaagctttg 2320
 acatgggagg agaaactctg ttgtctgtct gtcccatggg 2360
 gccgaaggac ttgggcaa atcttgtctgt gcaaaacctg 2400
 atcaggtcga ccgaggaact gaatatataa ctttcaggga 2440

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FIG. 1D

gtgagtcaag tggctccaga ggcagccaag atttttaccc 2480
caaatggagg gaatccaaat tgtttataac tgatgaagag 2520
gtgggtcccg aagagacaga gacagacact tttgatgccg 2560
caccgcagcc tgccagggaa gctgcctttg catcagactc 2600
tctaaggact ggaaggtcac gatcatctca gagcatttgt 2640
aaggcaggag aaagtacaga tgccctcagc ttgcctcatg 2680
tcaaactgaa ataa 2694

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FIG. 2A

KCNQ5 Protein Sequence

MKDVESGRGR	VLLNSAAARG	DGLLLLGT	ATLGGGGGGL	40
RESRRGKQGA	RMSLLGKPLS	YTSSQSCRN	VKYRRVQNYL	80
YNVLERPRGW	AFIYHAFVFL	LVFGCLILSV	FSTIPEHTKL	120
ASSCLLILEF	VMIVVFGLEF	IIRIWSAGCC	CRYRGWQGRL	160
RFARKPFCVI	DTIVLIASIA	VVSAKTQGNI	FATSALRSLR	200
FLQILRMVRM	DRRGGTWKLL	GSVVYAHSKE	LITAWYIGFL	240
VLIFSSFLVY	LVEKDANKEF	STYADALWWG	TITLTTIGYG	280
DKTPLTWLGR	LLSAGFALLG	ISFFALPAGI	LGSGFALKVQ	320
EQHRQKHFEK	RRNPAANLIQ	CVWRSYAADE	KSVSIATWKP	360
HLKALHTCSP	TKKEQGEASS	SQKLSFKERV	RMASPRGQSI	400
KSRQASVGDR	RSPSTDITAE	GSPTKVQKSW	SFNDRTRFRP	440
SLRLKSSQPK	PVIDADTALG	TDDVYDEKGC	QCDVSVEDLT	480
PPLKTVIRAI	RIMKFHVAKR	KFKETLRPYD	VKDVIEQYSA	520
GHLDMLCRIK	SLQTRVDQIL	GKGQITSDDK	SREKITAEHE	560
TTDDL SMLGR	VVKVEKQVQS	IESKLDCLLD	IYQQVLRKGS	600
ASALALASFQ	IPPFCEQTS	DYQSPVDSKD	LSGSAQN SGC	640
LSRSTSANIS	RGLQFILTPN	EFSAQTFYAL	SPTMHSQATQ	680
VPISQSDGSA	VAATNTIANQ	INTAPKPAAP	TTLQIPPLP	720
AIKHLPRPET	LHPNPAGLQE	SISDVTTCLV	ASKENVQVAQ	760
SNLTKDRSMR	KSFDMGGETL	LSVCPMVPKD	LGKSLSVQNL	800
IRSTEELNIQ	LSGSESSGSR	GSQDFYPKWR	ESKLFITDEE	840

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FIG. 2B

VGPEETETDT FDAAPQPARE AAFASDSLRT GRSRSSQSIC	880
KAGESTDALS LPHVKLK	897

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FIG. 3

Alternative Splice Exon 1

TGG	GGA	CAG	TGG	ACA	TTG	CGT
Trp	Gly	Gln	Trp	Thr	Leu	Arg

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FIG. 4A

3' UTR

gttcttcatt	ttctttccag	gcatagcagt	tcttttagcca	40
tacatatcat	tgcatgaact	atttcgaaag	cccttctaaa	80
aagttgaaat	tgcaagaatc	gggaagaaca	tgaaaggcag	120
tttataagcc	cgttaccttt	taattgcatg	aaaatgcatg	160
tttagggatg	gctaaaattc	caaggtgcat	cgacattaac	200
ccactcattt	agtaatgtac	cttgagttaa	aaagcctgag	240
aaaccaaaca	cagctaattgc	tatgggggtgt	atgaatatgt	280
caagtttagg	tcattttagaa	gatttgacac	tgtattttga	320
aattatgagt	aaacaccttc	aaatttcagg	catttctgct	360
ttgtgactaa	atacaaacta	cattttcaag	attaggccat	400
aatgtatatt	taaacacaat	ggctatcaac	agctgctaata	440
aaggatatcaa	ctaaagcaga	attgggggaat	aatagaaatg	480
gctgcttatt	tcaagatata	tttgccaacc	cattcctatt	520
cagtcatttt	attattaatg	taatttgaat	gtcaatttgt	560
gtgcttttgg	tgatttagcg	ctgtggcaag	caattttgca	600
catcattttc	atgttggtct	ttatgacaag	aatgtttctc	640
aattagaaaa	tgtgcaaata	atgaaattca	gggccagtga	680
ggcaaataga	ctatctgaca	tatttgactt	tatgaaaaca	720
tattgcctga	tggcagaatc	aactttataa	gtggtcaact	760
tctacacaag	cgtatgaaat	actggtcagt	agaacagcca	800

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FIG. 4B

ttgtgattgg actggtttct ctgcaatggc gccaacccca	840
ggcttgccaa tactgcctat gtaaagggca agtgtgagaa	880
gctattctca ttctgctgac atacaggtag gactatgggg	920
gatgggacat ttgagtggga ctgagatagg aaaggcttga	960
aaagaaccca gaaacaccac caggaagttg gcaaagtaaa	1000
agaaaatgac ttccccctca aaggggcaatg agagggagag	1040
aaacaaacca aaatagaaga actagacttt ttagaaaatg	1080
agtattgcta	1090

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FIG. 5A

Multiple Sequence Alignment of the KCNQ Channel Family Members

hskcnq4 ~~~~~MAEAPPRRLGLGPPPGDAPRAE.LVALTAVQSEQEAG.
hskcnq5 ~~~~~MKDVESGR.GRVLLNSAAARGDGLLLLGTRAATLGGG.
hskcnq2 ~~~~~MVQKSRNG.GVYPG..PSGEKKLKVGFGVGLDPG.AP.
hskcnq3 MGLKARRAAGAAGGGGDGGGGGGGAANPAGDAAAAGDEERKVGLAPGDVEQVTLALCA.
hskcnq1 ~~~~~MAAASSPPRAERKRWGRLPGARRGSAGLAKKCPFSLEAECCP

FIG. 5B

hskcnq4 ESSRMGIKDRIRMGSSQRRRTGPS.KQQLAPPTMPTSPSSSEQVGEATSPTKVQKSWSFNDR
hskcnq5 SSQKLSFKERVIRMASPRGQSIKS.RQ..ASVGDRRSPSTDITAEG.SP TKVQKSWSFNDR
hskcnq2 .SOKVSLKDRV.FSSPRGVAAKG.KGSPQAQTVRRSPSADQSLE.DSPSKVPKSWSFGDR
hskcnq3 SSQKLGLLDRVRLSNPRGSNTKG.KLF.....TPLNVDAIE.ESPSKEPKPVGLNNK
hskcnq1 PGEKMLTVPHITCDPPEERRLDHFSVDGYDSSVRKSET...LLEVSMPE.....

hskcnq4 TRFRASLRL....KPRTSABDA.PSEEVAEEKSYQCELTVDDEMPAVKTVIRSTRIRIKK
hskcnq5 TRFRPSLRLKSSQPKVIDADTALGTDDVYDEKGCQCDVSVEDLTPPLKTVIRATRIRIKK
hskcnq2 SRARQAFRIKGAASR.QNSEEASLPGEDIVDDKSCPCFVTEDETPGLKVSIRAVCMVR
hskcnq3 ERFRTAFRMKAYAFWQ..SSEDAGTGDPMAEDRGYGNDFPIEDMIPTLKAATRAVRITOE
hskcnq1 .HF...MRTNSFAEDLDLEGETLLT..PITH.....ISQREHHKATIKVIRRMQY

hskcnq4 LVAKRKFKETLRPYDVKDVIEWEQYSAGHLDMLGRIKSLQTRMDQIVGRG...PGDR.KARE
hskcnq5 HVAKRKFKETLRPYDVKDVIEWEQYSAGHLDMLCRIKSLQTRMDQILGKGQI.TSDK.KSRE
hskcnq2 LVSKRKFKESLRPYDVMVIEWEQYSAGHLDMLSRIKSLQSRMDQIVGRGPA.ITD..KDR.
hskcnq3 RLYKKFKETLRPYDVKDVIEWEQYSAGHLDMLSRIKYLQTRMDMIFTPGPP.STPKHKKSQ
hskcnq1 FVAKKKFQOARKPYDVRDVIEWEQYSOGLNLMVRIKELORRLDQSIGKPSLFISVSEKSKD

hskcnq4 KGDKG.....PSDAEVVDEISMMGRVVKVE..KQVQSIEHKLDLLLGFY
hskcnq5 ...KI.....TAEHETDLSMLGRVVKVE..KQVQSIESKLDCLLDIY
hskcnq2 ..TKG.....PAEAELPEDPSMMGRLGKVE..KQVLSMEKKLDLFLVNIY
hskcnq3 KGSFTFPSQQSPRNEPYVARPSTSEI.EDQSMMGKFVKVE..RQVQDMGKKLDLFLVDMH
hskcnq1 RGSNTIGARLNRVEDKVTQLDQRLALITD...MLHQLLSLHGGSTPGSGGPPREGGAHIT

hskcnq4 SRCLRSGT..SA.SLGAVQVPLFDPDITSYHSFVDEH..EDISVSAQTLS.ISRSVSTNM
hskcnq5 QQVLRKGS.SALALASFQIPPFCEQTSYQSPVDS..KDLGSAQNSGCLSRSTSANI
hskcnq2 MQ..RMGIP.PTETEAYFGAK..EPEPAPPYHSPEDS..RE...HVDRHGCIVKIVRSSS
hskcnq3 MQHMER.....LQVQVTEYYPTKGTSSPAEAEKKEDNRYSDLKTIICNYSETGP
hskcnq1 QPCGSGGSVDPELFLPSNTLPTYE.QLTVPRRGPDGGS~~~~~

hskcnq4 D~~~~~
hskcnq5 SRGLQFI..LTPNEFSAQTFYALSPTMHSQATQVPISQSDGSAVAATNTIANQINTAPKP
hskcnq2 STGQKNF..SAPPAAPP...VQCPPSTSWQPQSHPRQGHGTSPVGDHGS�VRIPPPPAH
hskcnq3 PEPPYSFHQVTIDKVSPYGFFAHDPVNLPRGGPSS.GKVQATPPSSATTYVERPTVLPIL
hskcnq1 ~~~~~~

hskcnq4 ~~~~~~
hskcnq5 AAPTTLQIPPLPAIKHLRPETLHPNPAGLQESISDVTTCLVASKENVQVAQSNLTKDR
hskcnq2 ERSLSAYGGGNRASMEFLRQEDTPGCRPPEGTLRDSDTISIPSVDEELERSFSFGFSIS
hskcnq3 TLLDSRVSCHSQADLQG.PYSDRISPRQRRSITRSDTPLSLMSVNHEELERSPSGFSIS
hskcnq1 ~~~~~~

hskcnq4 ~~~~~~
hskcnq5 SMRKSFDMGGETLLSVCPMPVK...DLGKSLSVQNLIRSTEELNIQLSGSESSGSRGSQ
hskcnq2 QSKENLDALNSCYAAVAPCAKVRPYIAEGESDTSDDLCTPCGPPPRSATGEGPFQDVGWA
hskcnq3 QDRDDYVFGPN...GGSSWMREKRYLAEGETDTDTPFTPSGSMPLSSTGDGIDS SVWTP
hskcnq1 ~~~~~~

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FIG. 5C

hskcnq4 ~~~~~
hskcnq5 DFYPKWRESKLFITDEEVGPETETDTFDAAPQPAREAAFASDSLRTGRSRSSQSICKAG
hskcnq2 GPRK~~~~~
hskcnq3 SNKPI~~~~~
hskcnq1 ~~~~~

hskcnq4 ~~~~~
hskcnq5 ESTDALSLPHVKLK
hskcnq2 ~~~~~
hskcnq3 ~~~~~
hskcnq1 ~~~~~

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FIG. 6A

Human RNA Master Blot

TABLE 1

	1	2	3	4	5	6	7	8
A	whole brain	amygdala	Caudate nucleus	cerebellum	cerebral cortex	frontal lobe	hippocampus	medulla oblongata
B	occipital pole	putamen	substantia nigra	temporal lobe	thalamus	Subthalamic nucleus	spinal cord	
C	heart	aorta	Skeletal muscle	colon	bladder	uterus	prostate	stomach
D	testis	ovary	pancreas	pituitary gland	adrenal gland	thyroid gland	salivary gland	mammary gland
E	kidney	liver	small intestine	spleen	thymus	peripheral leukocyte	lymph node	bone marrow
F	Appendix	lung	trachea	placenta				
G	fetal brain	fetal heart	fetal kidney	fetal liver	fetal spleen	fetal thymus	fetal lung	
H	yeast total RNA	yeast tRNA	<i>E. coli</i> rRNA	<i>E. coli</i> DNA	Poly r(A)	human C ₀ t DNA	human DNA	human DNA

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FIG. 6B

	1	2	3	4	5	6	7	8
A	•		•		•	•	•	
B	•	•		•				
C			•					
D								
E								
F								
G								
H				•				

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FIG. 7

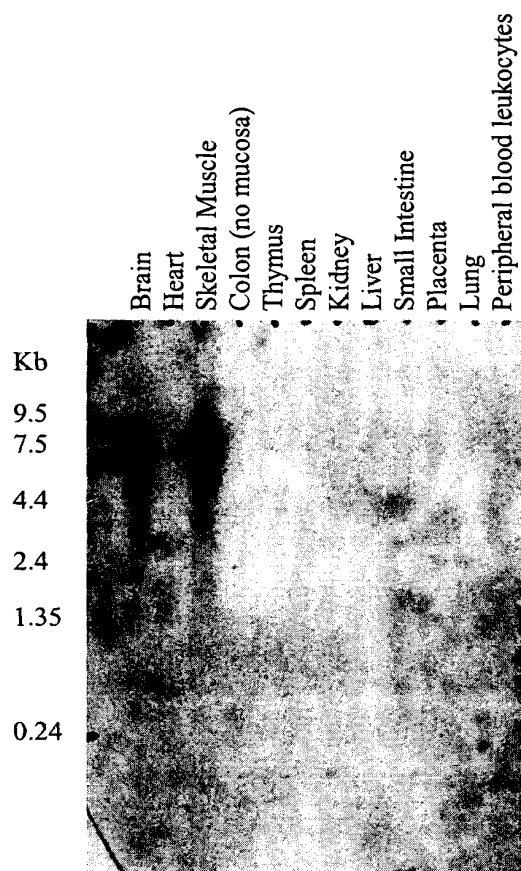
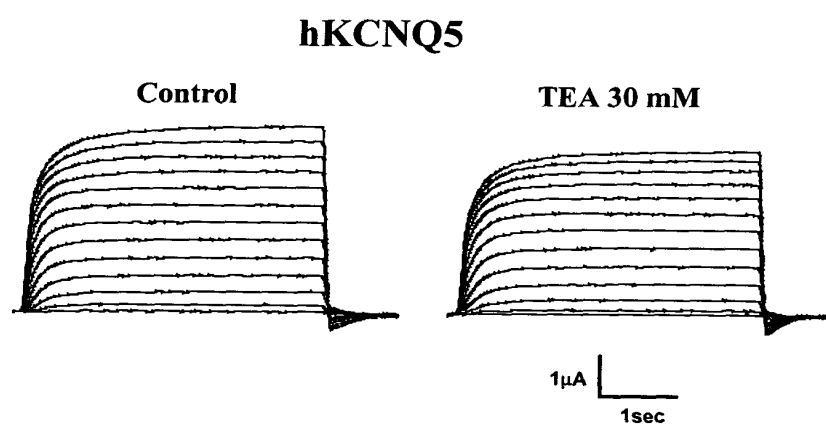


FIG. 8



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Antisense



FIG. 9A

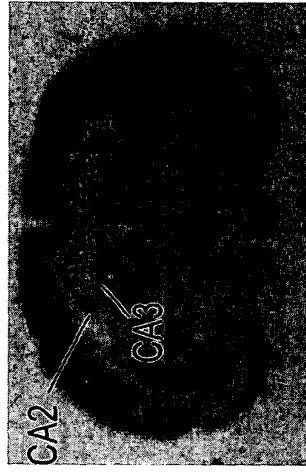


FIG. 9B



FIG. 9C

Sense



FIG. 9D

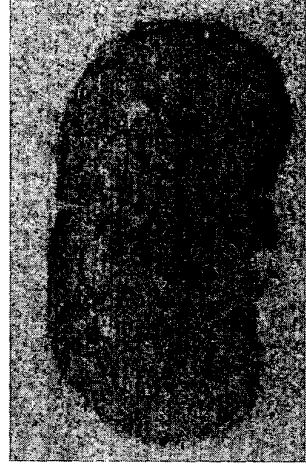


FIG. 9E



FIG. 9F

FIG. 10A

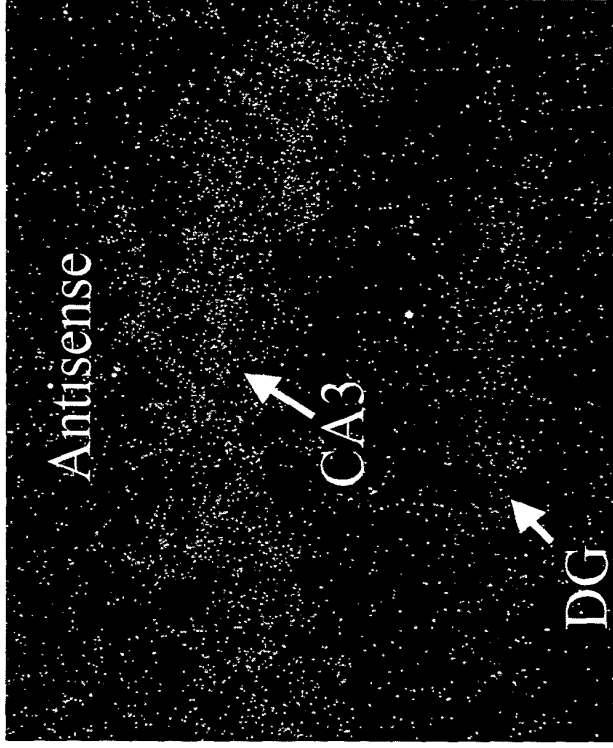


FIG. 10B

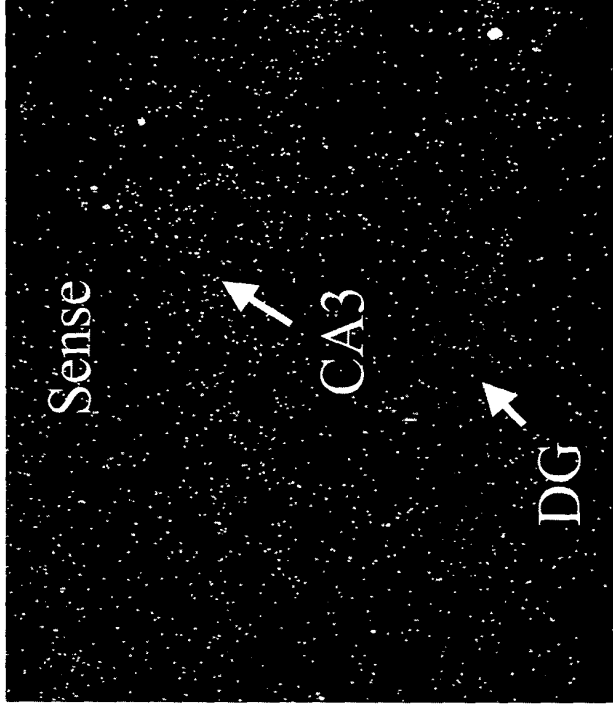


FIG. 11

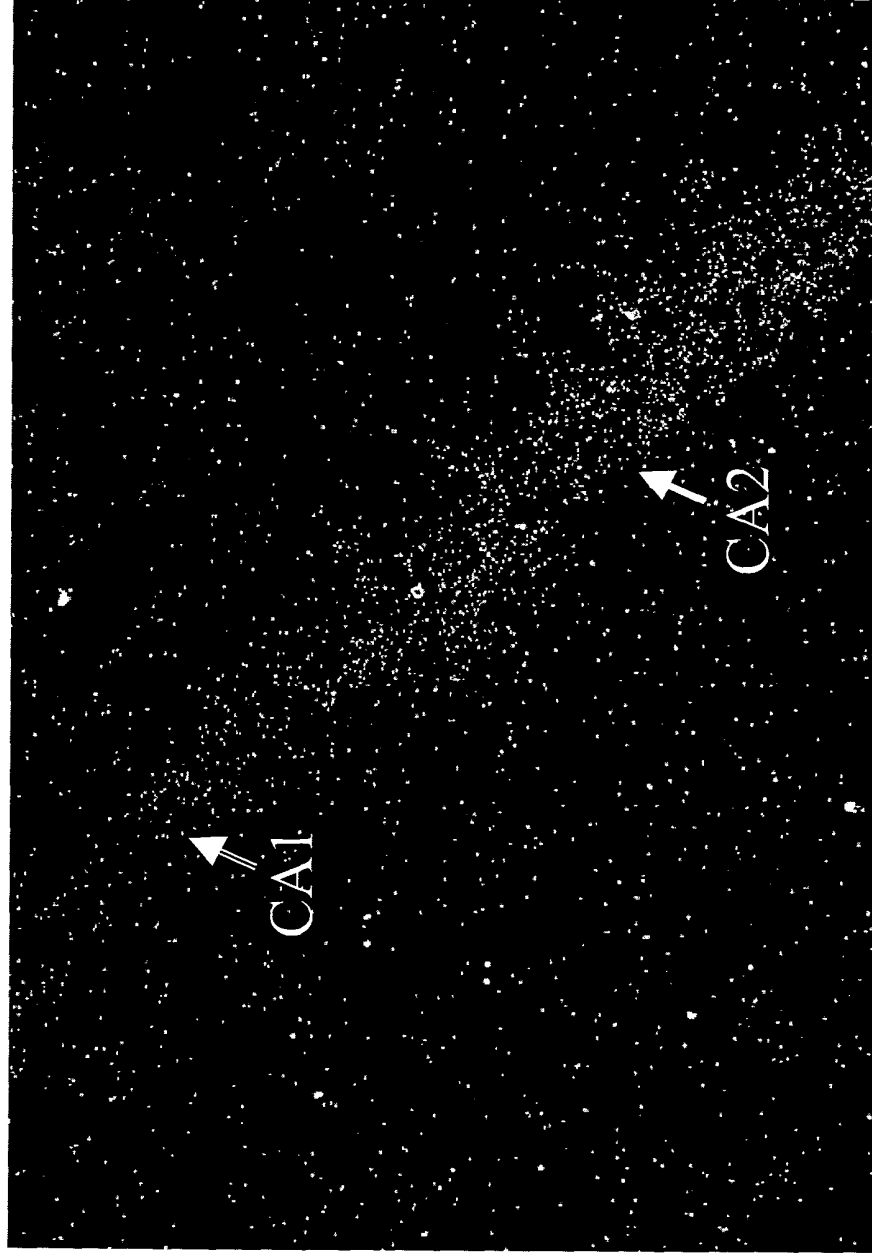


FIG. 12A

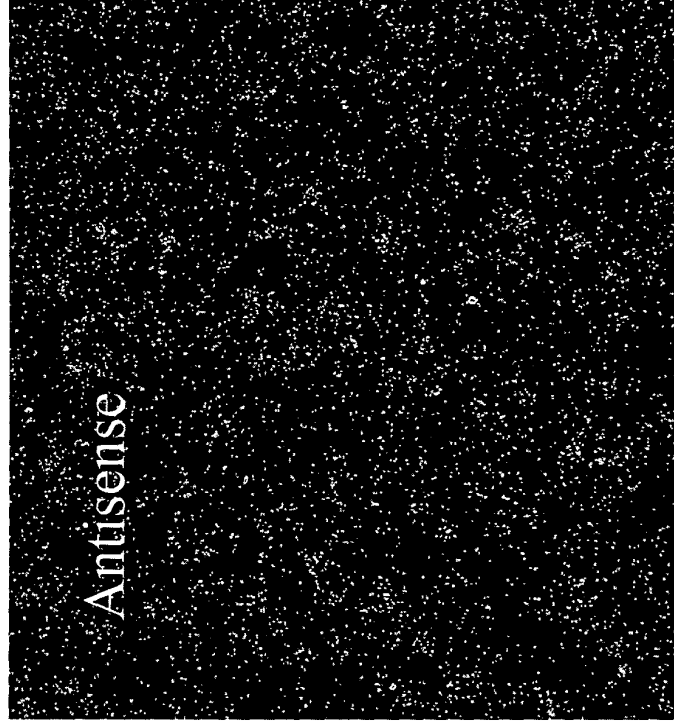


FIG. 12B

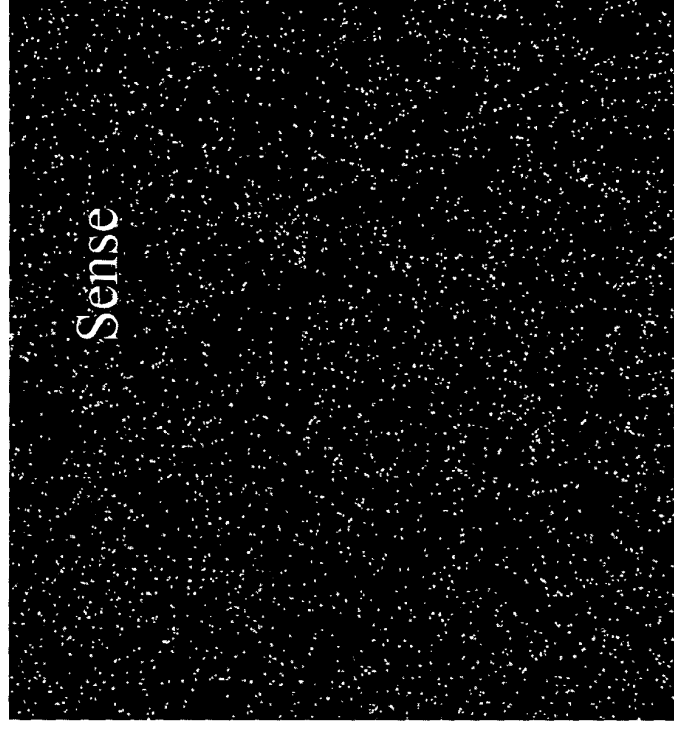


FIG. 13A

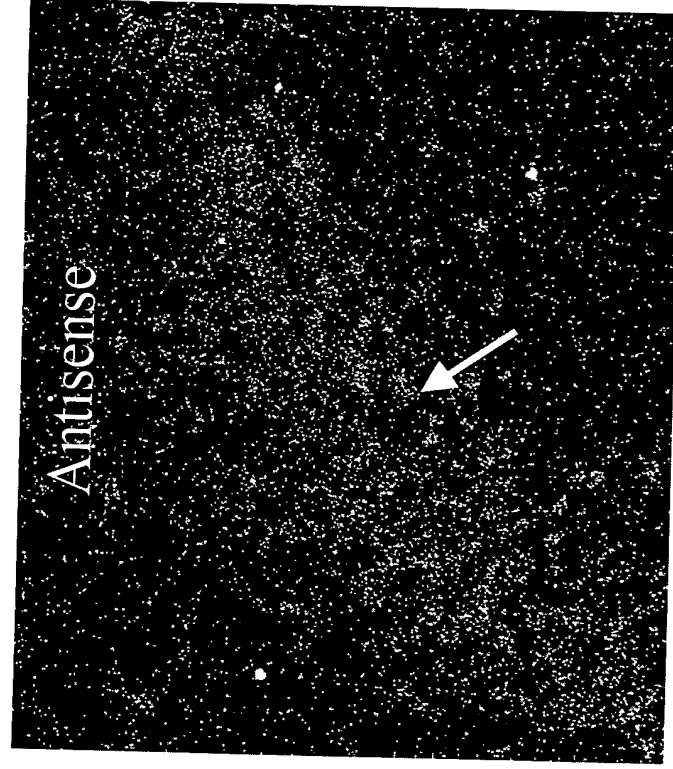


FIG. 13B

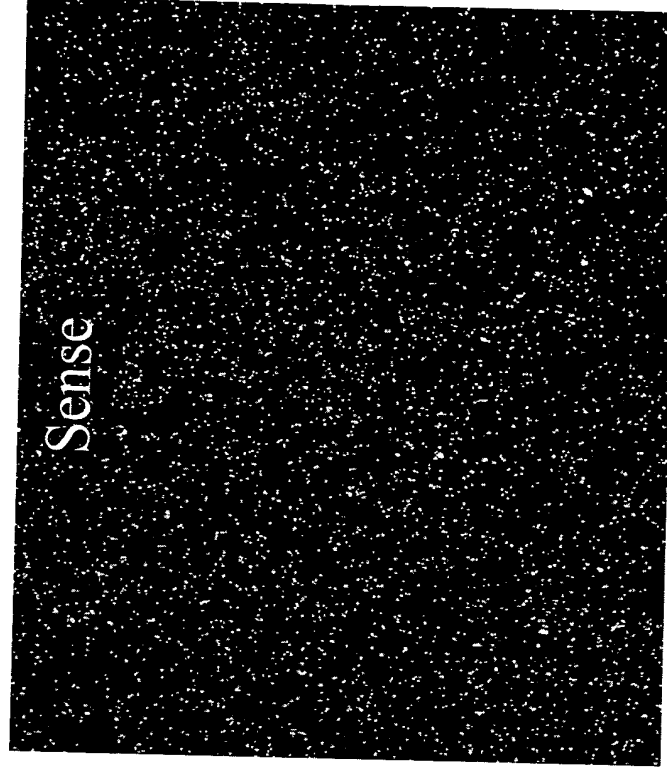


FIG. 14A

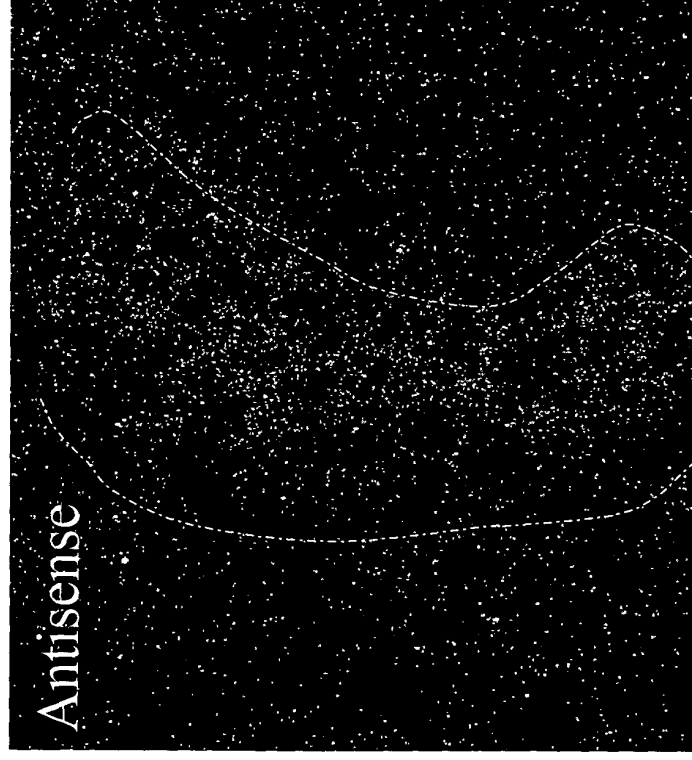


FIG. 14B

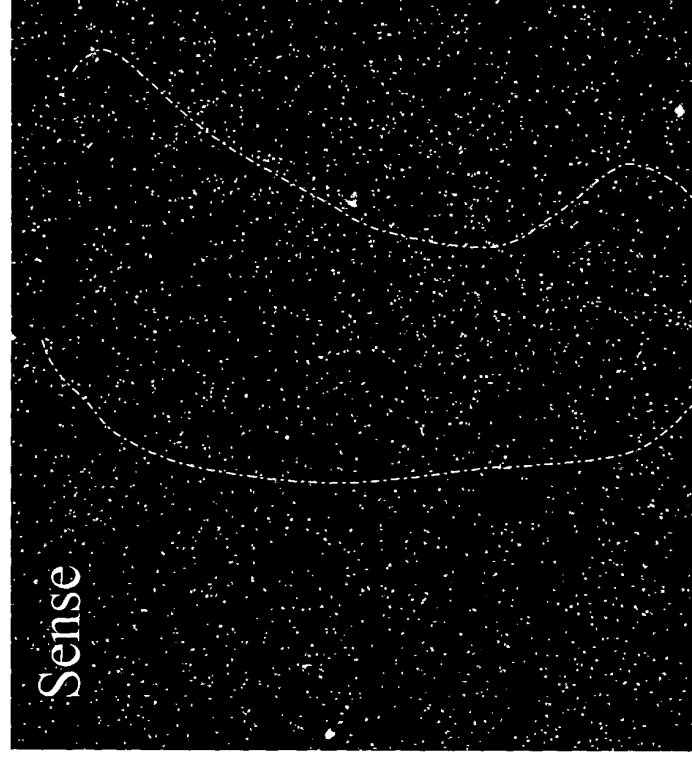


FIG. 15A

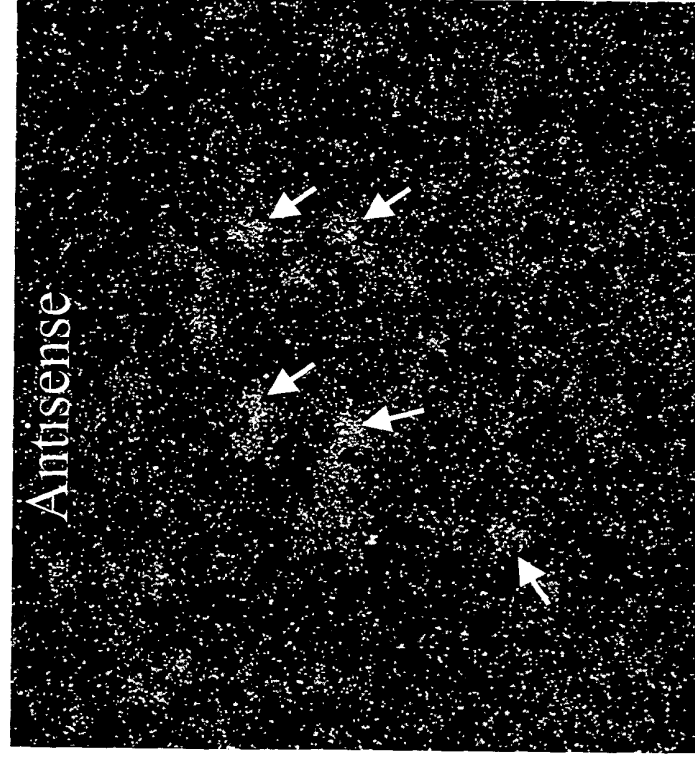


FIG. 15B

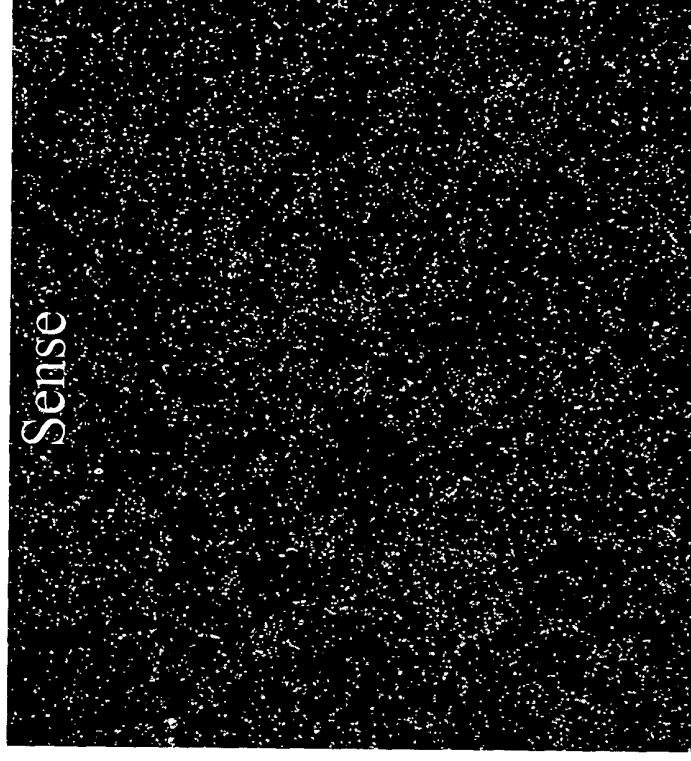


FIG. 16A

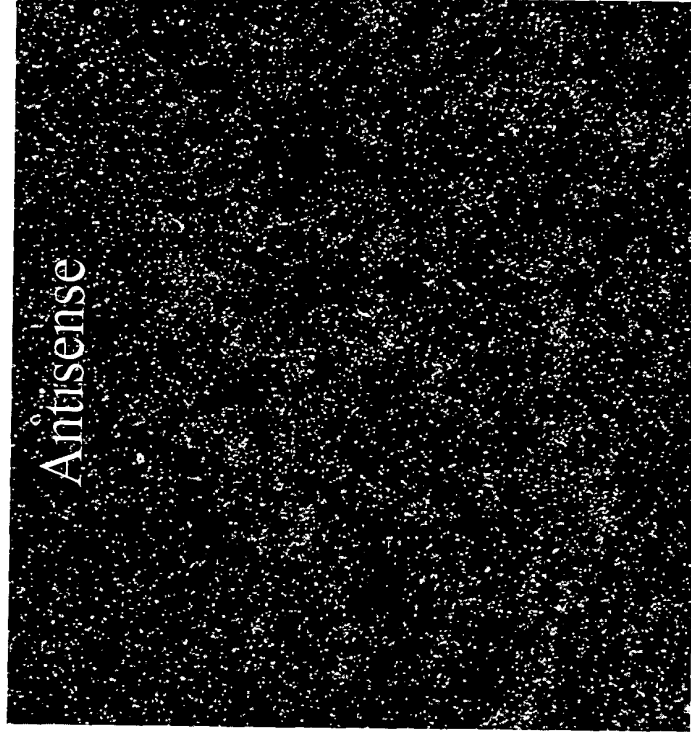


FIG. 16B

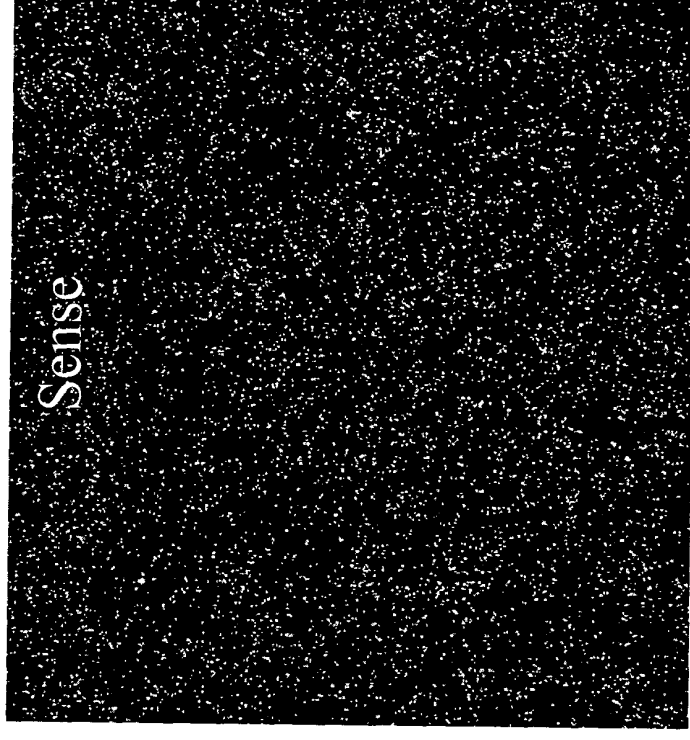


FIG. 17A

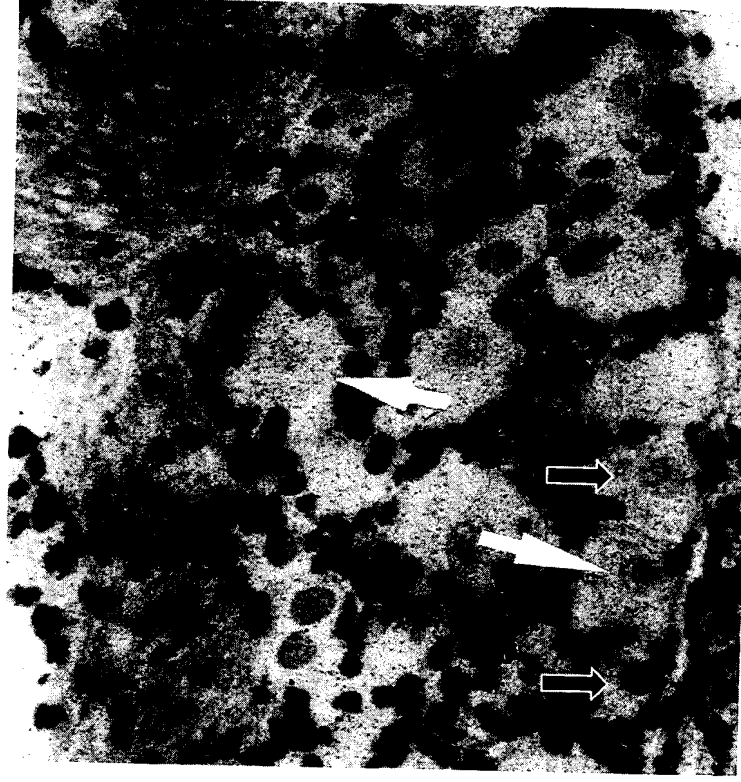


FIG. 17B

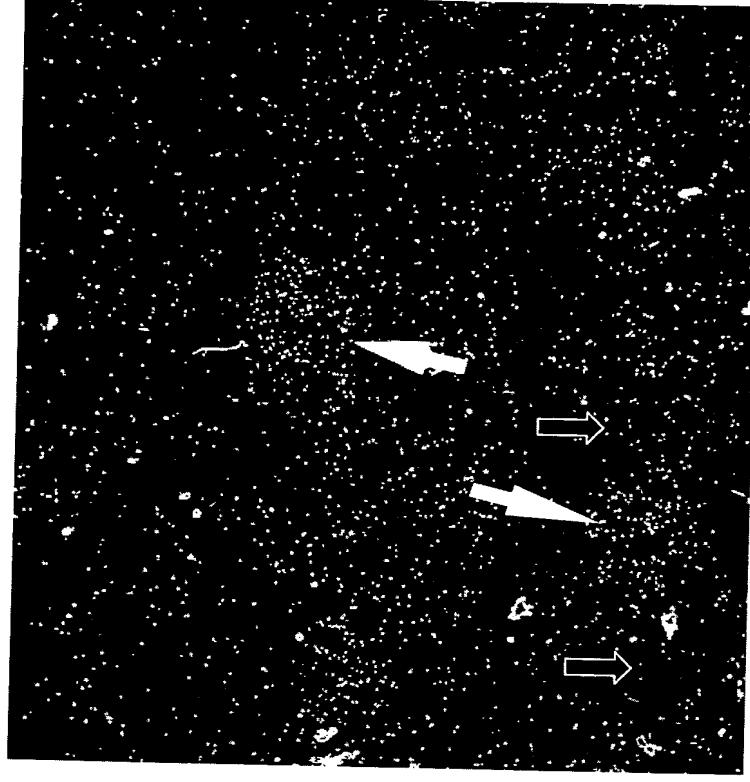


FIG. 18A

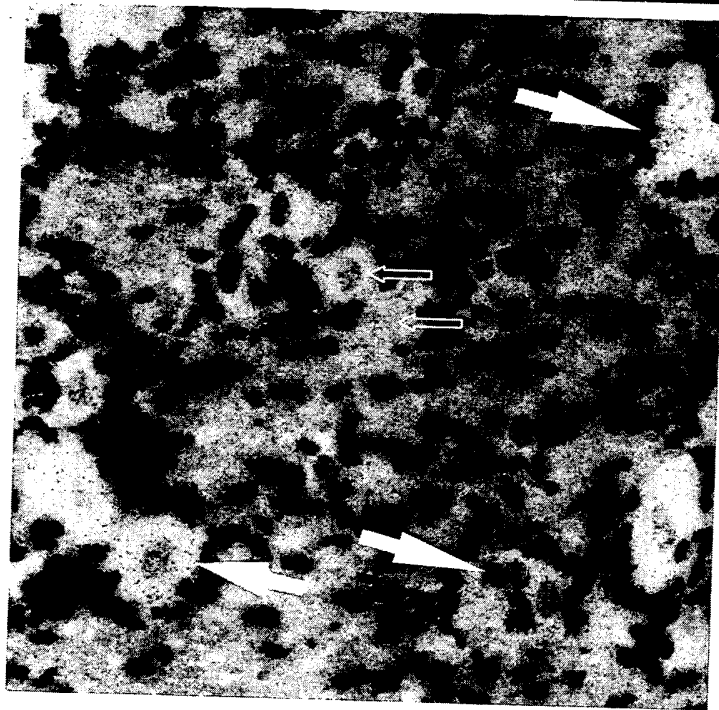


FIG. 18B

